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RESEARCH ARTICLE

Wearable derived cardiovascular responses to stressors in free-living conditions

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Abstract

Stress contributes to the progression of many diseases. Despite stress' contribution towards disease, few methods for continuously measuring stress exist. We investigated if continuously measured cardiovascular signals from a wearable device can be used as markers of stress. Using wearable technology (WHOOP Inc, Boston, MA) that continuously measures and calculates heart rate (HR) and heart rate variability (root-mean-square of successive differences; HRV), we assessed duration and magnitude of deviations in HR and HRV around the time of a run (from 23665 runs) or high-stress work (from 8928 high-stress work events) in free-living conditions. HR and HRV were assessed only when participants were motionless (HR_{motionless}). Runs were grouped into light, moderate, and vigorous runs to determine dose response relationships. When examining HR_{motionless} and HRV throughout the day, we found that these metrics display circadian rhythms; therefore, we normalized HR_{motionless} and HRV measures for each participant relative to the time of day. Relative to the period within 30 minutes leading up to a run, HR_{motionless} is elevated for up to 180-210 minutes following a moderate or vigorous run (P<0.05) and is unchanged or reduced following a light run. HRV is reduced for at least 300 minutes following a moderate or vigorous run (P<0.05) and is unchanged during a light run. Relative to the period within 30 minutes leading up to high-stress work, HR_{motionless} is elevated during and for up to 30 minutes following highstress work. HRV tends to be lower during high-stress work (P = 0.06) and is significantly lower 90-300 minutes after the end of the activity (P<0.05). These results demonstrate that wearables can quantify stressful events, which may be used to provide feedback to help individuals manage stress.

Introduction

Stress has been implicated as a contributing factor to numerous diseases, including cardiovascular disease, depression, and certain types of cancers [1, 2]. The number of stressful events, the degree to which someone responds to a stressful event, and the time it takes to recover from a stressful event have been found to be associated with mortality and poor mental and physical health [3, 4]. Despite the links between stress and disease, few methods exist that allow people to objectively quantify the amount of stress caused by an event; therefore, many

these authors are articulated in the 'author contributions' section.

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individuals lack the knowledge to avoid or properly manage stressful events. To appropriately manage stress, methods that quantify the impact and duration of stressful events in real-time need to be developed.

In controlled laboratory settings, stress is typically gauged with surveys, cardiovascular measures, or blood/saliva tests. However, each of these methods are limited in their ability to be used for widespread use: surveys require conscientious feedback that may not be possible during or right after a stressful event; cardiovascular measures are classically acquired using leads for electrocardiograms and sphygmomanometers for measuring blood pressure, which require bulky equipment and specialized knowledge to use; and blood or saliva tests require specialized equipment and trained technicians to measure the concentration of stress hormones. Translating these laboratory-grade stress examinations into tests that are easier to perform and interpret may promote greater adoption of stress tracking.

Certain wearable devices continuously measure cardiovascular parameters using photoplethysmography and provide a unique opportunity to investigate the magnitude of stressful events on cardiovascular parameters in real-time. In particular, HR and HRV have the potential to be surrogate markers of stress, whereby both HR or HRV can vary in response to the degree of sympathetic or parasympathetic input [5–7]. Recent research has investigated the ability of wearable-derived metrics to detect or measure stressful events, like infections and vaccines, when aggregated over a long periods of time [8–10]. In this study, we aimed to leverage the continuous monitoring provided by the WHOOP strap (WHOOP Inc, Boston, MA) to determine the magnitude of stressful events on cardiovascular metrics in free-living conditions.

Methods

Data collection

Since data were not identifiable and were stored on a secure server, this study was deemed exempt from Institutional Review Board (IRB) oversight by Advarra's IRB (Columbia, MD). Participants were excluded from the study if they were under the age of 21 and if they indicated a gender other than male or female. A wrist-worn device (WHOOP versions 3.0 and 4.0; Boston, MA) that continuously collects heart rate, three-axis accelerometer, temperature, and three-axis gyroscope data was used to calculate cardiovascular metrics and sleep and wake times, which have been validated elsewhere [11]. Running and high-stress work activities are logged via manual input from a participant or auto-detected by the WHOOP analytics platform. Sleep and wake onset are autodetected. To be considered for this analysis, a user must have had at least 25 days of data over a 28-day span and no more than 28 days of data were used for each participant.

Data calculation

Resting heart rate and heart rate variability. Heart rate ($HR_{motionless}$) and heart rate variability (HRV) were calculated during periods that lacked motion detected by the accelerometer. Moving-block sub-sampling was used to calculate $HR_{motionless}$ and HRV. The step size was 30 seconds and utilized a 5-minute block of photoplethysmography-derived beat-to-beat intervals and accelerometry data to calculate $HR_{motionless}$ and HRV in the absence of motion artifacts. Heart rate variability was calculated as the root-mean-square of successive differences. $HR_{motionless}$ and HRV derived from the WHOOP device have been validated against gold-standard measures from electrocardiograms [11].

Relative fractional differences. To calculate the relative fractional difference for resting heart rate and heart rate variability throughout the day, we first calculated the median resting

heart rate and heart rate variability for each hour of the day over a 28-day period for each participant. We then divided the median daily $HR_{motionless}$ and HRV by each hour's median $HR_{motionless}$ and HRV value to determine the relative fractional difference of each hour's $HR_{motionless}$ and HRV from the median daily $HR_{motionless}$ and HRV within each user.

Impact of an activity. $HR_{motionless}$ and HRV were aggregated over the duration of the activity or over 30-minute intervals before the start of or following an activity. Aggregated $HR_{motionless}$ and HRVs were then subtracted by the median hourly $HR_{motionless}$ and HRV for that time of day for a given user. Lastly, we subtract the post-activity normalized $HR_{motionless}$ and HRV values from the pre-activity normalized $HR_{motionless}$ and HRV values. Cardiovascular parameters were not calculated during the running activity due to motion artifacts which prevent the reliable acquisition of the photoplethysmography data [12]. To determine whether a running load was light, moderate, or vigorous, we utilized a proprietary algorithm that scores a run using the amount of time spent in a cardiovascular zone based on an individual's heart rate reserve.

Statistical analysis. Data analyses were conducted in either Python (version 3.8.12) or R (version 4.2.0). Descriptive statistics are provided as frequencies and percentages for categorical variables or means and 95% confidence intervals for continuous variables. To assess circadian rhythms in $HR_{motionless}$ and HRV, we fit splines to linear models to determine the non-linear relationships. To determine if $HR_{motionless}$ and HRV deviated from baseline relative to the period before a run or high-stress work, we first modeled the data using linear mixed-effect models and then utilized Dunnet's test as a post-hoc test that directly compared the time intervals following an activity to the time before an activity. To determine the difference in $HR_{motionless}$ and HRV between running loads, we calculated areas under the curve using the trapezoidal method [13] (treating 0 as the baseline) for each running load group and compared these areas under the curve using Tukey's test. Significance was set as P < 0.05.

Results

Diurnal fluctuations in HR_{motionless} and HRV

To better understand the diurnal variation that occurs throughout the day, we analyzed normalized hour-to-hour fluctuations in $HR_{motionless}$ and HRV throughout the day for 974 individuals (Table 1) aggregated within a given clock hour or hours to and from sleep or wake.

Table 1. Demographics of participants for circadian analysis.

	Overall (N = 974)				
Gender					
female	273 (28.0%)				
male	701 (72.0%)				
Age					
Mean (CI)	36.849 (36.196, 37.503)				
Height					
Mean (CI)	1.766 (1.760, 1.772)				
Weight					
Mean (CI)	80.850 (79.848, 81.852)				
BMI					
Mean (CI)	25.818 (25.560, 26.075)				

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Within a given clock hour (Fig 1A and 1B), HR_{motionless} decreases from time point 0.5 to 4.5 (β = -0.014, P<0.001), increases from time point 4.5 to 13.5 (β = 0.025, P<0.001), negligibly decreases from time point 13.5 to 19.5 (β = -0.004, P<0.001), and decreases from time point 19.5 to 23.5 (β = -0.026, P<0.001). For HRV within a given clock hour (Fig 1B), we find that HRV increases from time point 0.5 to 6.5 (β = 0.042, P<0.001) and time point 6.5 to 11.5 (β = 0.077, P<0.001), and decreases from time point 11.5 to 17.5 (β = -0.027, P<0.001) and 17.5 to 23.5 (β = -0.073, P<0.001).

We next modeled fluctuations in $HR_{motionless}$ and HRV leading up to and following sleep, with negative numbers reflecting hours until sleep onset and positive numbers reflecting hours after sleep (Fig 1C and 1D). $HR_{motionless}$ increases from time point -14.5 to -9.5 (β = 0.014, P<0.001), negligibly decreases from -9.5 to -3.5 (β = -0.002, P<0.001), decreases from -3.5 to 6.5 (β = -0.024, P<0.001), and negligibly increases from 6.5 to 8.5 (β = 0.022, P<0.001). HRV increases from time point -14.5 to -11.5 (β = 0.068, P<0.001), decreases from -11.5 to -0.5 (β = -0.036, P<0.001) and -0.5 to 0.5 (β = -0.284, P<0.001), and increases from 0.5 to 8.5 (β = 0.031, P<0.001).

Lastly, we modeled fluctuations in $HR_{motionless}$ and HRV leading up to and following wake, with negative numbers representing hours until wake and positive numbers representing hours after wake (Fig 1E and 1F). $HR_{motionless}$ decreases from time point -8.5 to -0.5 (β = -0.009, P<0.001), increases from -0.5 to 2.5 (β = 0.073, P<0.001), and negligibly decreases from 2.5 to 12.5 (β = -0.001, P<0.001) and from 12.5 to 14.5 (β = -0.022, P<0.001). HRV increases from time point -8.5 to -0.5 (β = 0.03, P<0.001) and from -0.5 to 1.5 (β = 0.193, P<0.001), negligibly increases from 1.5 to 4.5 (β = 0.019, P<0.001), and decreases from 4.5 to 14.5 (β = -0.038, P<0.001).

Direct and residual effects of physical and psychological stressors on HR_{motionless} and HRV

To characterize the effects of a physical stressor on cardiovascular parameters, we examined HR $_{\rm motionless}$ and HRV starting from 30 minutes leading up to and 5 hours following 23665 running activities (Table 2). Furthermore, we aimed to determine if we could detect a dose effect of running on HR $_{\rm motionless}$ and HRV and grouped runs into light, moderate, and vigorous loads based on time spent in cardiovascular zones. Relative to the period before the run, HR $_{\rm motionless}$ remains elevated after a moderate or vigorous run for at least 180–210 minutes (P<0.05) and is unchanged or decreases after a light run (Fig 2A). Using the area under the curve to compare the relative change in HR $_{\rm motionless}$ post-run between running loads we find that, compared to light runs, both moderate and vigorous runs led to greater elevations in HR $_{\rm motionless}$ (P<0.05; Fig 2B). Relative to the period before the run, HRV remains depressed after a moderate and vigorous run for at least 270–300 minutes (P<0.05) and is unchanged after a light run (Fig 2C). Using the area under the curve to compare the relative change in HRV post-run between running loads, we find that vigorous runs lead to larger decreases in HRV than moderate to vigorous runs (P<0.05; Fig 2D).

To characterize the effects of a psychological stress on cardiovascular parameters, we examined $HR_{motionless}$ and HRV starting from 30 minutes leading up to and 300 minutes following 8928 high-stress work periods (Table 3). We find that $HR_{motionless}$ is elevated during and for up to 30-minutes after high-stress work (P<0.05; Fig 3A) and after thirty-minutes is either unchanged or depressed (P>0.05; Fig 3A). HRV trended towards being lower during high-stress work (P = 0.06; Fig 3B) and was significantly lower 90–300 minutes after the event (P<0.05; Fig 3B). In summary, these results suggest that $HR_{motionless}$ and HRV from wearables may be used as surrogate markers of physical or psychologically stressful events.

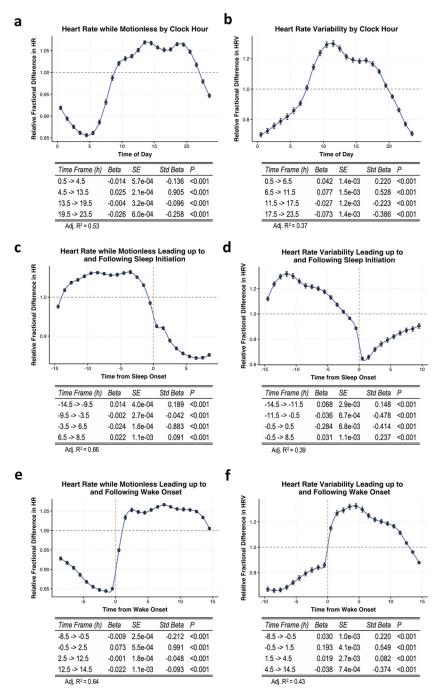


Fig 1. Diurnal variations in $HR_{motionless}$ and HRV. Data are aggregated for each hour over 28 days for each participant and expressed as the relative fractional difference from the average throughout the day. (a) $HR_{motionless}$ relative to time of day. (b) HRV relative to time of day. (c) $HR_{motionless}$ leading up to and following sleep onset. (d) HRV leading up to and following sleep onset. (e) $HR_{motionless}$ leading up to and following wake onset. (f) HRV leading up to and following wake onset. Tables below each figure represent the results from the linear spline models. Data are represented by means \pm 95% CI.

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Table 2. Demographics of running populations and descriptive statistics of associated runs.

	Light (N = 463)	Moderate (N = 1834)	Vigorous (N = 1097)		
Gender					
Female	221 (47.7%)	833 (45.4%)	502 (45.8%)		
Male	242 (52.3%)	1001 (54.6%)	595 (54.2%)		
Age					
Mean (CI)	37.404 (36.414, 38.394)	36.437 (35.969, 36.906)	35.486 (34.893, 36.079)		
Height (m)					
Mean (CI)	1.732 (1.723, 1.742)	1.734 (1.730, 1.739)	1.732 (1.726, 1.737)		
Weight (kg)					
Mean (CI)	74.983 (73.462, 76.505)	73.834 (73.166, 74.501)	72.237 (71.443, 73.032)		
ВМІ					
Mean (CI)	24.793 (24.420, 25.167)	24.402 (24.241, 24.564)	23.954 (23.765, 24.142)		
Number of Runs					
Count	1640.000	17085.000	4940.000		
Number of Runs Logged per Person					
Mean (CI)	3.542 (3.098, 3.986)	9.316 (9.029, 9.602)	4.503 (4.245, 4.761)		
Max Heart Rate (BPM)					
Mean (CI)	146.536 (144.519, 148.552)	174.751 (174.330, 175.171)	183.068 (182.567, 183.569)		
Average Heart Rate (BPM)					
Mean (CI)	116.283 (114.278, 118.288)	144.803 (144.321, 145.284)	154.626 (154.072, 155.180)		
Duration (min)					
Mean (CI)	26.239 (21.662, 30.816)	43.746 (42.975, 44.516)	83.818 (80.490, 87.146)		

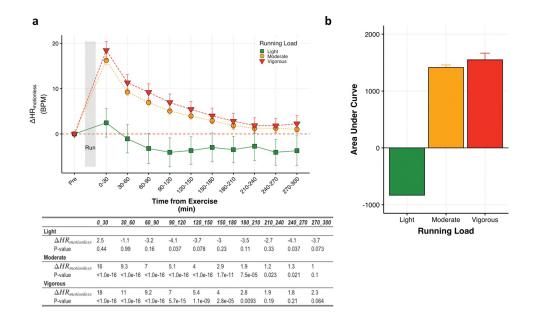
Heart rate includes measures taken in the presence of motion.

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Discussion

The response to a stressor can be measured via reactivity—how much the system deviates from baseline—and recovery—how long it takes the system to return to baseline [14]—and both reactivity and recovery can be quantified using cardiovascular parameters [15]. Improving awareness into how the body responds to stress, termed interoception, may lead to healthier stress responses [16]; therefore, easily-accessible methods to quantify the cardiovascular impact of stressors, which may improve interoception, may help individuals manage their stress. In this investigation, we collated self-reported physically and psychologically stressful events with objective measures from a wearable that continuously measures cardiovascular parameters (WHOOP Inc, Boston, MA). We find that individuals wearing WHOOP display circadian rhythms in their HR_{motionless} and HRV, and that deviations from an individual's circadian rhythm can be used to gauge the magnitude of a stressful event based on HR_{motionless} and HRV.

A key finding from this study is that $HR_{motionless}$ and HRV exhibit a circadian rhythm that aligns with the time a person falls asleep or wakes up. This finding is novel as this study is the first to investigate patterns of $HR_{motionless}$ and HRV relative to when someone falls asleep or wakes up. Using clock time to assess circadian rhythms, prior research indicates that HR is elevated during day time and decreases leading up to and throughout the night, whereas HRV increases throughout sleep, peaks in the early morning, and subsequently decreases throughout the day [17, 18]. Since HR and HRV exhibit circadian rhythms, measures that rely on these cardiovascular parameters may need to be corrected for the time of day when the measure was taken.



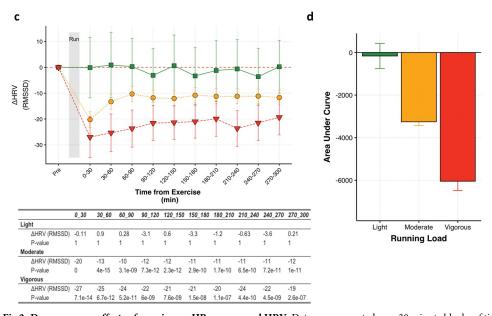


Fig 2. Dose-response effects of running on HR_{motionless} and HRV. Data are aggregated over 30 minute blocks of time leading up to and following a run, and are normalized to time of day and the thirty minutes leading up to the run ("Pre"). (a) Change in HR_{motionless} (BPM) relative to the period before a run. The table reflects the absolute differences and P-values for HR_{motionless} following a run, grouped by running load. P-values were determined using Dunnet's test, treating the period before a run ("Pre") as the control group. (b) Area under the curve for HR_{motionless} following a run. P-values were determined using Tukey's test with comparisons made across all groups. (c) Change in HRV (RMSSD) relative to the period before a run. The table reflects the absolute differences and P-values for HRV following a run, grouped by running load. P-values were determined using Dunnet's test, treating the period before a run ("Pre") as the control group. (d) Area under the curve for HRV following a run. P-values were determined using Tukey's test with comparisons made across all groups. Data are represented by means ± 95% CI.

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Table 3. Demographics of high-stress work population and descriptive statistics of associated high-stress work events.

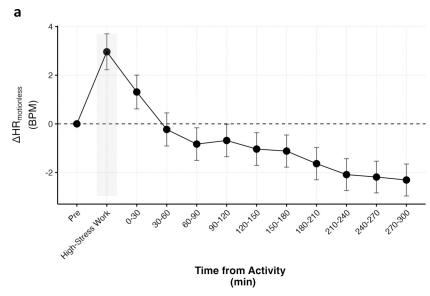
	Overall (N = 1778)				
Gender					
Female	536 (30.1%)				
Male	1242 (69.9%)				
Age					
Mean (CI)	36.916 (36.339, 37.493)				
Height (m)					
Mean (CI)	1.756 (1.751, 1.760)				
Weight (kg)					
Mean (CI)	80.338 (79.527, 81.148)				
BMI					
Mean (CI)	25.916 (25.704, 26.127)				
Number of High-Stress Works					
Count	8928.000				
Number of High-Stress Works Logged per Person					
Mean (CI)	5.021 (4.625, 5.418)				
Max Heart Rate (BPM)					
Mean (CI)	149.561 (148.576, 150.545)				
Average Heart Rate (BPM)					
Mean (CI)	113.630 (112.586, 114.674)				
Duration (min)					
Mean (CI)	114.792 (108.945, 120.638)				

Heart rate includes measures taken in the presence of motion.

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Correcting for the time of day when HR_{motionless} and HRV were measured, we sought to determine how different types of stressors might impact cardiovascular dynamics in real-time conditions. High-intensity physical stressors, like exercise, can elevate HR and reduce HRV for many hours after the physical stressor has been removed due to persistent activation of the sympathetic nervous system and inhibition of the parasympathetic nervous system [19, 20]. Conversely, low-intensity exercise has been shown to reduce heart rate for up to 24 hours following the exercise bout, possibly due to an acute reduction in total and regional vascular resistance [21, 22]. We find that we can replicate these laboratory-based findings in real-world conditions by examining the impact of different running loads on cardiovascular dynamics post-exercise. These findings indicate that data derived from wearable devices can be used to assess the impact of an exercise session on the cardiovascular system, which may be useful in determining the efficacy of the exercise and assessing short-term recovery from exercise. Moreover, these results may also have health implications as the rate of cardiovascular recovery from exercise has been shown to be a predictor of mortality [23].

We also investigated how cardiovascular dynamics change in response to self-perceived high-stress work, which may consist of a psychological stress, physical stress, or a combination of the two. Psychological stress impacts cardiovascular metrics in a similar manner as physical stress—by increasing HR and reducing HRV during and following a stressful event [24, 25]. In this investigation, we find that HR_{motionless} is elevated during and immediately after high-stress work, whereas HRV tends to be reduced during and following 90 minutes after high-stress work. One plausible explanation for the 90-minute lag observed with a stressor-induced



	During	0_30	30_60	60_90	90_120	120_150	150_180	180_210	210_240	240_270	270_300
$\Delta HR_{motionless}$	3	1.3	-0.23	-0.83	-0.68	-1	-1.1	-1.6	-2.1	-2.2	-2.3
P-value	3e-14	2e-03	1e+00	1e-01	3e-01	2e-02	9e-03	1e-05	3e-09	5e-10	2e-10

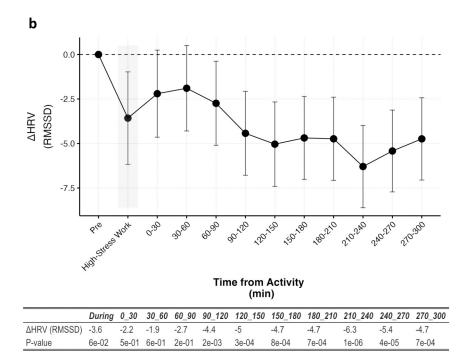


Fig 3. Effects of high-stress work on HR_{motionless} **and HRV**. Data are aggregated over 30 minute blocks of time leading up to, during, and following high-stress work, and are normalized to time of day and the thirty minutes leading up to the high-stress work. (a) Change in HR $_{motionless}$ (BPM) relative to the period before high-stress work. (b) Change in HRV (RMSSD) relative to the period before high-stress work. The table reflects the absolute differences and P-values for HR $_{motionless}$ or HRV following a high-stress work. P-values were determined using Dunnet's test, treating the period before a high-stress work ("Pre") as the control group. Data are represented by means \pm 95% CI.

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reduction in HRV following high-stress work is that this lagged-response may be due to an immediate relief of the stressor being removed (i.e., HRV returning to near pre-stressor levels) followed by a rumination period [26]; however, this hypothesis is only speculative and requires further research to validate.

In summary, the results from this study indicate that $HR_{motionless}$ and HRV exhibit a circadian rhythm that can be used as a baseline to detect deviations in $HR_{motionless}$ and HRV caused by stressful events. These results have practical applications as they demonstrate a novel method of normalization to detect deviations in stress-related metrics and provide evidence that detecting stress can be done passively and outside of clinical settings. An interesting future direction for research will be to assess if these wearable-derived cardiovascular markers of stress can predict the onset of certain diseases [27]. Ultimately, these findings may be useful in detecting stressful events, increasing interoception, and improving responses to stressors.

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Investigation: David M. Presby.Methodology: David M. Presby.Visualization: David M. Presby.

Writing – original draft: David M. Presby, Summer R. Jasinski, Emily R. Capodilupo. Writing – review & editing: David M. Presby, Summer R. Jasinski, Emily R. Capodilupo.

References

- Cohen S, Janicki-Deverts D, Miller GE. Psychological stress and disease. JAMA. 2007; 298: 1685– 1687. https://doi.org/10.1001/jama.298.14.1685 PMID: 17925521
- Eckerling A, Ricon-Becker I, Sorski L, Sandbank E, Ben-Eliyahu S. Stress and cancer: mechanisms, significance and future directions. Nat Rev Cancer. 2021; 21: 767–785. https://doi.org/10.1038/s41568-021-00395-5 PMID: 34508247
- Crosswell AD, Lockwood KG. Best practices for stress measurement: How to measure psychological stress in health research. Health Psychol Open. 2020; 7: 2055102920933072. https://doi.org/10.1177/ 2055102920933072 PMID: 32704379
- Chida Y, Steptoe A. Greater cardiovascular responses to laboratory mental stress are associated with poor subsequent cardiovascular risk status: a meta-analysis of prospective evidence. Hypertension. 2010; 55: 1026–1032. https://doi.org/10.1161/HYPERTENSIONAHA.109.146621 PMID: 20194301
- Kannankeril PJ, Le FK, Kadish AH, Goldberger JJ. Parasympathetic effects on heart rate recovery after exercise. J Investig Med. 2004; 52: 394–401. https://doi.org/10.1136/jim-52-06-34 PMID: 15612453
- Goldberger JJ, Challapalli S, Tung R, Parker MA, Kadish AH. Relationship of heart rate variability to parasympathetic effect. Circulation. 2001; 103: 1977–1983. https://doi.org/10.1161/01.cir.103.15.1977 PMID: 11306527
- Trimmel K, Sacha J, Huikuri HV. Heart Rate Variability: Clinical Applications and Interaction between HRV and Heart Rate. Frontiers Media SA; 2015.

- Alavi A, Bogu GK, Wang M, Rangan ES, Brooks AW, Wang Q, et al. Real-time alerting system for COVID-19 and other stress events using wearable data. Nat Med. 2022; 28: 175–184. https://doi.org/ 10.1038/s41591-021-01593-2 PMID: 34845389
- Presby DM, Capodilupo ER. Biometrics from a wearable device reveal temporary effects of COVID-19 vaccines on cardiovascular, respiratory, and sleep physiology. J Appl Physiol. 2022; 132: 448–458. https://doi.org/10.1152/japplphysiol.00420.2021 PMID: 35019761
- Quer G, Radin JM, Gadaleta M, Baca-Motes K, Ariniello L, Ramos E, et al. Wearable sensor data and self-reported symptoms for COVID-19 detection. Nat Med. 2021; 27: 73–77. https://doi.org/10.1038/ s41591-020-1123-x PMID: 33122860
- Miller DJ, Sargent C, Roach GD. A Validation of Six Wearable Devices for Estimating Sleep, Heart Rate and Heart Rate Variability in Healthy Adults. Sensors. 2022; 22: 6317. https://doi.org/10.3390/ s22166317 PMID: 36016077
- Seok D, Lee S, Kim M, Cho J, Kim C. Motion Artifact Removal Techniques for Wearable EEG and PPG Sensor Systems. Frontiers in Electronics. 2021;2. https://doi.org/10.3389/felec.2021.685513
- Signorell A, Aho K, Alfons A, Anderegg N, Aragon T, Arppe A, et al. DescTools: Descriptive Tools Analysis. In: https://cran.r-project.org/web/packages/desctools/index.html. 2022.
- Miller R, Wojtyniak J-G, Weckesser LJ, Alexander NC, Engert V, Lehr T. How to disentangle psychobiological stress reactivity and recovery: A comparison of model-based and non-compartmental analyses of cortisol concentrations. Psychoneuroendocrinology. 2018; 90: 194–210. https://doi.org/10.1016/j.psyneuen.2017.12.019 PMID: 29370954
- Clay JM, Parker MO. The role of stress-reactivity, stress-recovery and risky decision-making in psychosocial stress-induced alcohol consumption in social drinkers. Psychopharmacology. 2018; 235: 3243–3257. https://doi.org/10.1007/s00213-018-5027-0 PMID: 30209533
- Quadt L, Garfinkel SN, Mulcahy JS, Larsson DE, Silva M, Jones A-M, et al. Interoceptive training to target anxiety in autistic adults (ADIE): A single-center, superiority randomized controlled trial. EClinical-Medicine. 2021; 39: 101042. https://doi.org/10.1016/j.eclinm.2021.101042 PMID: 34401684
- Vandewalle G, Middleton B, Rajaratnam SMW, Stone BM, Thorleifsdottir B, Arendt J, et al. Robust circadian rhythm in heart rate and its variability: influence of exogenous melatonin and photoperiod. J Sleep Res. 2007; 16: 148–155. https://doi.org/10.1111/j.1365-2869.2007.00581.x PMID: 17542944
- Natarajan A, Pantelopoulos A, Emir-Farinas H, Natarajan P. Heart rate variability with photoplethysmography in 8 million individuals: a cross-sectional study. Lancet Digit Health. 2020; 2: e650–e657. https://doi.org/10.1016/S2589-7500(20)30246-6 PMID: 33328029
- Seiler S, Haugen O, Kuffel E. Autonomic recovery after exercise in trained athletes: intensity and duration effects. Med Sci Sports Exerc. 2007; 39: 1366–1373. https://doi.org/10.1249/mss. 0b013e318060f17d PMID: 17762370
- Terziotti P, Schena F, Gulli G, Cevese A. Post-exercise recovery of autonomic cardiovascular control: a study by spectrum and cross-spectrum analysis in humans. Eur J Appl Physiol. 2001; 84: 187–194. https://doi.org/10.1007/s004210170003 PMID: 11320634
- Quinn TJ. Twenty-four hour, ambulatory blood pressure responses following acute exercise: impact of
 exercise intensity. J Hum Hypertens. 2000; 14: 547–553. https://doi.org/10.1038/sj.jhh.1001106 PMID:
 10980585
- Rueckert PA, Slane PR, Lillis DL, Hanson P. Hemodynamic patterns and duration of post-dynamic exercise hypotension in hypertensive humans. Med Sci Sports Exerc. 1996; 28: 24–32. https://doi.org/10.1097/00005768-199601000-00010 PMID: 8775351
- Cole CR, Blackstone EH, Pashkow FJ, Snader CE, Lauer MS. Heart-rate recovery immediately after exercise as a predictor of mortality. N Engl J Med. 1999; 341: 1351–1357. https://doi.org/10.1056/NEJM199910283411804 PMID: 10536127
- Mohammadi A, Emamgoli A, Shirinkalam M, Meftahi GH, Yagoobi K, Hatef B. The persistent effect of acute psychosocial stress on heart rate variability. Egypt Heart J. 2019; 71: 18. https://doi.org/10.1186/s43044-019-0009-z PMID: 31659513
- Waugh CE, Panage S, Mendes WB, Gotlib IH. Cardiovascular and affective recovery from anticipatory threat. Biol Psychol. 2010; 84: 169–175. https://doi.org/10.1016/j.biopsycho.2010.01.010 PMID: 20096747
- Ottaviani C, Shapiro D, Davydov DM, Goldstein IB, Mills PJ. The autonomic phenotype of rumination. Int J Psychophysiol. 2009; 72: 267–275. https://doi.org/10.1016/j.ijpsycho.2008.12.014 PMID: 19272312
- Abbad Ur Rehman H, Lin C-Y, Mushtaq Z, Su S-F. Performance analysis of machine learning algorithms for thyroid disease. Arab J Sci Eng. 2021; 46: 9437–9449.